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IN THE UNITED STATES PATENTS AND TRADEMARKS OFFICE

APPLICANTS: Andrew VAILLANT et al.

ASSIGNEE: REPLICOR INC.

SERIAL NUMBER: 10/661,088

TITLE: ANTIVIRAL OLIGONUCLEOTIDES TARGETING HBV

FILING DATE: September 12, 2003

ART UNIT: 1648

EXAMINER: Bo, PENG

INFORMATION DISCLOSURE STATEMENT UNDER 37 CFR § 1.97 and 1.98

Commissioner for Patents PO Box 1450 Alexandria, Virginia 22313-1450

Sir:

Submitted herewith on a PTO/SB/08A/B form is a listing of documents known to Applicants in order to comply with Applicants' duty of disclosure pursuant to 37 CFR § 1.56. A copy of each listed document is being submitted to comply with the provisions of 37 CFR §§ 1.97-1.98.

The submission of any document herewith, which is not a statutory bar, is not intended as an admission that such document constitutes prior art against the claims of the

present application or that such document is considered material to patentability as defined in 05/10/2006 HASFRY1 00000060 195113 10661088

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Serial No. 10/661,088

37 CFR § 1.56(b). Applicants do not waive any rights to take any action which would be

appropriate to antedate or otherwise remove as a competent reference any document which is

determined to be a prima facie prior art reference against the claims of the present

application.

RELEVANCE OF EACH DOCUMENT

Applicants respectfully request that any listed document be considered by the

Examiner and be made of record in the present application and that an initialed copy of Form

PTO/SB/08A/B be returned in accordance with MPEP § 609.

The Commissioner is hereby authorized to withdraw the fees in the amount of

\$180.00 for the submission of an Information Disclosure Statement from Deposit Account

No. 19-5113 as well as any additional fees which may be required regarding this application

under 37 CFR §§ 1.16-1.17, or credit any overpayment, to Deposit Account No. 19-5113.

Respectfully submitted,

May 4, 2006

Christian Cawthorn, Reg. No. 47,352

Agent of record

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Encl. Form PTO/SB/08A/B

Non-Patent References in pdf format

PTO/SB/08A/B 08-03)

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INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(use as many sheets as necessary)

Sheet 1 of 13

Complete if Known						
Application Number	10/661,088					
Filing Date	September 12, 2003					
First Named Inventor	Andrew Vaillant et al.					
Art Unit	1648					
Examiner Name	Bo PENG					
Attorney Docket Number	16051-9US CC					

	U.S. PATENT DOCUMENTS							
Examiner	Cite	Document Number	Publication Date	Name of Patentee or	Pages, Columns, Lines Where Relevant Passages or Relevant			
Initials* No.	No.'	Number – Kind Code ² (if known)	MM-DD-YYYY	Applicant of Cited Document	Figures Appear			
		US-5,563,050						
		US-6,506,559						
		US-5,023,252						
		US-5,580,859						
		US-4,806,463						
		US-5,248,670						
		US-5,591,720						
		US-5,952,490						
		US-5,998,602						
		US-6,184,369						
		US-5,264,423						
		US-5,276,019						
		US-6,316,190						
		US-5,218,103						
		US-5,684,148						
		US-5,452,496						
		US-5,278,302						
		US-5,750,666						
		US-5,602,244						
		US-5,508,270						

	FOREIGN PATENT DOCUMENTS								
Examiner	Cite	Foreign Patent Document	Publication Date	Name of Patentee or	Pages, Columns, Lines	T			
Initials*	No.1	Country Code3 - Number4 - Kind Code5 (if known)	MM-DD-YYYY	Applicant Of Cited Document	Where Relevant Passages or Relevant Figures Appear	T ⁶			
		WO 03/02903		, , , =					
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		WO 94/02499				1			
		WO 94/26764				1			
		WO 97/13499				1			

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			U.S. PATENT D	OCUMENTS	
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		US-4,469,863			i i i i i i i i i i i i i i i i i i i
-		US-5,610,289			
		US-5,256,775			
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		US-5,476,925			
	•	US-5,023,243			
		US-5,130,302			
		US-5,177,198			
		US-5,378,825			
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		US-5,264,562	<u> </u>		
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		US-5,770,713			
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		US-4,534,899			
•		US-5,705,188			
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	FOREIGN PATENT DOCUMENTS								
Examiner Initials*	Cite	Foreign Patent Document	Publication Date	Name of Patentee or	Pages, Columns, Lines	Τ			
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		WO 90/04384							
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		WO 94/20073				+			
		WO 96/10391				_			
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		US-5,213,804			
		US-5,225,212			
		US-5,540,935			
		US-5,556,948			
		US-6,316,190			
	-	US-5,264,221			
		US-5,665,710			
		US-4,806,463			
		US-5,248,670			
		US-5,695,979			
		US-5,591,623			
		US-5,514,788			
		US-5,652,355			
		US-6,143,881			
		US-6,346,614			
		US-5,591,721			
	_	US-6,608,035			
		US-3,687,808			
		US-5,625,050			

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		US-6,573,099				
		US-6,013,240				
		US-5,837,449				
		US-6,177,246				
i		US-5,670,634				
		US-2003/232435				
		US-2003/0119019 A1				
		US-2003/0162225 A1				
					 	

FOREIGN PATENT DOCUMENTS						
Cite	Foreign Patent Document	Publication Date	Name of Patentee or	Pages, Columns, Lines	T	
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		NON PATENT LITERATURE DOCUMENTS	
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T²
		ADJOU et al., "A novel generation of heparan sulfate mimetics for the treatment of prion diseases", 2003, J. Gen. Virol. 84:2595-2603.	
		AGRAWAL, "Importance of nucleotide sequence and chemical modifications of antisense oligonucleotides", 1999, <i>Biochim. Biophys. Acta</i> 1489:53-68.	
		AGRAWAL and KANDIMALLA, "Antisense therapeutics: is it as simple as complementary base recognition?", 2000, <i>Mol. Med. Today</i> 6:72-81.	
		AKHTAR et al., "The delivery of antisense therapeutics", 2000, Advanced Drug Delivery Reviews 44:3-21.	
		ALLAKHVERDI et al., "Inhibition of Antigen-induced Eosinophilia and Airway Hyperresponsiveness by Antisense Oligonucleotides Directed against the Common β Chain of IL-3, IL-5, GM-CSF Receptors in a Rat Model of Allergic Asthma", 2002, Am. J. Respir. Crit. Care Med. 165:1015-1021.	
		ANDREOLA et al., "DNA aptamers selected against the HIV-1 RNase H display in vitro antiviral activity", 2001, Biochemistry, 40:10087-10094.	
		BAKER et al., "2-O-(2-Methoxy)ethyl-modified Anti-intercellular Adhesion Molecule 1 (ICAM-1) Oligonucleotides Selectively Increase the ICAM-1 mRNA Level and Inhibit Formation of the ICAM-1 Translation Initiation Complex in Human Umbilical Vein Endothelial Cells", 1997, J. Biol. Chem. 272 (18):11994-12000.	
		BALL et al., "Clinical Potential of Respirable Antisense Oligonucleotides (RASONs) in Asthma", 2003, Am. J. Pharmacogenomics 3 (2):97-106.	
		BANKS <i>et al.</i> , "Delivery across the Blood-Brain Barrier of Antisense Directed against Amyloid β: Reversal of Learning and Memory Deficits in Mice Overexpressing Amyloid Precursor Protein", 2001, <i>J. Pharmacol. Exp. Ther.</i> 297 (3):1113-1121.	
		BARDOS et al., "Structure-Activity Relationships and Mode of Action of 5-Mercapto-Substituted Oligo- and Polynucleotides as Antitemplates Inhibiting Replication of Human Immunodeficiency Virus Type 1", 1992, Antimicrob. Agents and Chemother. 36 (1):108-114.	

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		BARRET et al., "Evaluation of Quinacrine Treatment for Prion Diseases", 2003, J. of Virol. 77 (15):8462-8469.	
		BATE et al., "Squalestatin Cures Prion-infected Neurons and Protects Against Prion Neurotoxicity", 2004, J. of Biol. Chem. 279 (15):14983-14990.	
		BOUSSIF et al., "A versatile Vector for Gene and Oligonucleotide Transfer into Cells in Culture and in vivo: Polyethylenimine", 1995, <i>Proc. Natl. Acad. Sci. USA.</i> 92 (16):7297-7301.	
		BRIGGER et al., "Poly(ethylene glycol)-Coated Hexadecylcyanoacrylate Nanospheres Display a Combined Effect for Brain Tumor Targeting", 2002, <i>J. Pharmacol. Exp. Ther.</i> 303 (3):928-936.	
		CASPER, "Discovery of a Novel Target for Potential Cancer Therapy", Joint Bayer Science Forum – ACS November 24 th Dinner Meeting. 2003, The FILTERPAPER, Andy Edelbrock Bayer Corporation, page 3 (abstract).	
	-	CAUGHEY et al., "Sulfated Polyanion Inhibition of Scrapie-Associated PrP Accumulation in Cultured Cells", 1993, J. Virol. 67 (2):643-650.	
		CHEN et al., "Antisense Oligonucleotides Demonstrate a Dominant Role of c-Ki-RAS Proteins in Regulating the Proliferation of Diploid Human Fibroblasts", 1996, J. Biol. Chem. 271 (45):28259-28265.	
		CHEONG et al., "Structure of influenza virus panhandle RNA studied by NMR spectroscopy and molecular modeling", 1999, Nuc. Acids. Res. 27 (5): 1392-1397.	
		CHIANG et al., "Antisense Oligonucleotides Inhibit Intercellular Adhesion Molecule 1 Expression by Two Distinct Mechanisms", 1991, <i>J. Biol. Chem.</i> 266 (27):18162-18171.	
		CIOFFI et al., "Selective Inhibition of A-Raf and C-Raf mRNA Expression by Antisense Oligodeoxynucleotides in Rat Vascular Smooth Muscle Cells: Role of A-Raf and C-Raf in Serum-Induced Proliferation", 1997, Mol. Pharmacol. 51:383-389.	

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STA	ATEMENT	BY AP	PLICANT	First Named Inventor	Andrew Vaillant et al.	
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		NON PATENT LITERATURE DOCUMENTS	
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T ²
		CROOKE et al., "In Vitro Toxicological Evaluation of ISIS 1082, a Phosphorothioate Oligonucleotide Inhibitor of Herpes Simplex Virus", 1992, Antimicrob. Agents Chemother. 36 (3):527-532.	
		DASS, "Vehicles for oligonucleotide delivery to tumours", 2002, Journal of Pharmacy and Pharmacology 54:3-27.	
		DASS, "Liposome-Mediated Delivery of Oligodeoxynuclotides In Vivo", 2002, <i>Drug Delivery</i> , 9:169-180.	
		DHEUR et al., "Polyethylenimine but Not Cationic Lipid Improves Antisense Activity of 3'-Capped Phosphodiester Oligonucleotides", 1999, Antisense & Nucleic Acid Drug Development, 9:515-525.	
		DOH-URA et al., "Treatment of Transmissible Spongiform Encephalopathy by Intraventricular Drug Infusion in Animal Models", 2004, <i>J. Virol.</i> 78 (10):4999-5006.	
		ELBASHIR et al., "RNA interference is mediated by 21- and 22-nucleotide RNAs", 2001, Genes & Development 15:188-200.	
		ELBASHIR et al., "Duplexes of 21-nucleotide RNAs mediate RNA interference in cultured mammalian cells", 2001, Nature 411:494-498.	
		FINOTTO et al., "Local administration of antisense phosphorothioate oligonucleotides to the c-kit ligand, stem cell factor, suppresses airway inflammation and IL-4 production in a murine model of asthma", 2001, J. Allergy Clin. Immunol. 107 (2):279-286.	
		FISET et al., "Modulation of allergic response in nasal mucosa by antisense oligodeoxynucleotides for IL-4", 2003, J. Allergy Clin. Immunol. 111 (3):580-586.	
		GARRETT et al., "In vivo use of oligonucleotides to inhibit choroidal neovascularisation in the eye", 2001, J. Gene Med. 3:373-383.	

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		GORLACH et al., "Antisense repression in Cryptococcus neoformans as a laboratory tool and potential antifungal strategy", 2002, Microbiology 148:213-219.	
		GRIGORIEV et al., "Effects of the polyene antibiotic derivative MS-8209 on the astrocyte lysosomal system of scrapie-infected hamsters", 2002, <i>J. Mol. Neurosci.</i> 18:271-281.	
		HARBOTH et al., "Identification of essential genes in cultured mammalian cells using small interfering RNAs", 2001, Journal of Cell Science 114 (24) 4557-4565.	
		HORVATH et al., "Potent inhibition of HIV-1 entry by (s ⁴ dU) ₃₅ ", 2005, Virology 334:214-223.	
		HUGHES et al., "The cellular delivery of antisense oligonucleotides and ribozymes", 2001, Drug Discovery Today. 6 (6):303-315.	
		HUWYLER et al., "Brain drug delivery of small molecules using immunoliposomes", 1996, Proc. Natl. Acad. Sci. USA 93:14164-14169.	
		INOUE et al., "Synthesis and properties of novel nucleic acid probes", 1985, Symposium Series – Nucleic Acids Research No. 16:165-168.	
		INOUE et al., "Sequence-dependent hydrolysis of RNA using modified oligonucleotide splints and RNase H", 1987, FEBS Letters 215 (2):327-330.	
		JAASKELAINEN et al., "A lipid carrier with a membrane active component and a small complex size are required for efficient cellular delivery of anti-sense phosphorothioate oligonucleotides", 2000, European Journal of Pharmaceutical Sciences 10:187-193.	
		KANAGARATNAM et al., "Malaria merozoite surface protein antisense oligodeoxynucleotides lack antisense activity but function as polyanions to inhibit red cell invasion", 1998, Int. J. Biochem. Cell Biol. 30:979-985.	
		KOCISKO et al., "New Inhibitors of Scrapie-Associated Prion Protein Formation in a Library of 2,000 Drugs and Natural Products", 2003, J. Virol. 77 (19):10288-10294.	

Examiner	_	Date	52	
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		KOSTER et al., "Emerging therapeutic agents for transmissible spongiform encephalopathies: a review", 2003, J. Vet Pharmacol. Ther. 26:315-26.	
		KURRECK, "Antisense technologies. Improvement through novel chemical modifications", 2003, Eur. J. Biochem. 270:1628-1644.	
		KUWASAKI et al., "Inhibition of human immunodeficiency virus 1 replication in vitro by a self-stabilized oligonucleotide with 2'-O-methyl-guanosine-uridine quadruplex motifs", 2003, J. Antimicrob. Chemother. 51:813-819.	
		LAU et al., "Abrogation of c-Raf expression induces apoptosis in tumor cells", 1998, Oncogene 16:1899-1902.	
		LEUNG and SHAH, in: Controlled Release of drugs: Polymers and aggregate systems, Rosoff, M., Ed., 1989, VCH Publishers, New York, pages 185-215.	
		LUO et al., "Inhibition of influenza viral polymerases by minimal viral RNA decoys", 1997, J. Gen. Virol. 78: 2329-2333.	
		de MERGNY et al., "Kinetics and thermodynamics of i-DNA formation: phosphodiester versus modified oligodeoxynucleotides", 1998, Nucleic Acids Res. 26 (21): 4797-4803.	
		de MONBRISON et al., "Introducing antisense oligonucleotides into Pneumocystis carinii", 2002, J. Microbiol. Methods 50:211-213.	
		MONIA et al., "Evaluation of 2'-Modified Oligonucleotides Containing 2'-Deoxy Gaps as Antisense Inhibitors of Gene Expression", 1993, <i>J. Biol. Chem.</i> 268 (19) :14514-14522.	
		MORASSUTTI et al., "Effect of phosphorothioate modifications on the ability of GTn oligodeoxynucleotides to specifically recognize single-stranded DNA-binding proteins and to affect human cancer cellular growth" 1999, Biochimie 81:1115-1122.	

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Sheet	10	of	13	Attorney Docket Number	16051-9US CC	

		NON PATENT LITERATURE DOCUMENTS	
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		MOU and GRAY, "The high binding affinity of phosphorothioate-modified oligomers for Ff gene 5 protein is moderated by the addition of C-5 propyne or 2'-O-methyl modifications", 2002, <i>Nucleic Acids Res.</i> 30 (3):749-758.	
		NAKAJIMA et al., "Results of Quinacrine Administration to Patients with Creutzfeldt-Jakob Disease", 2004, Dement. Geriatr. Cogn. Disord. 17:158-163.	
		NOGUCHI et al., "Remarkable induction of apoptosis in cancer cells by a novel cationic liposome complexed with a bcl-2 antisense oligonucleotide", 2003, Journal of Controlled Release 88:313-320.	
		NOONPAKDEE et al., "Inhibition of Plasmodium falciparum proliferation in vitro by antisense oligodeoxynucleotides against malarial topoisomerase II", 2003, Biochem. and Biophys. Res. Commun. 302:659-664.	
		O'BRIEN et al., "Antisense BCR-ABL Oligomers Cause Non-Specific Inhibition of Chronic Myeloid Leukemia Cell Lines", 1994, Leukemia 8 (12):2156-2162.	
		OMORI et al., "Targeting of post-ischemic cerebral endothelium in rat by liposomes bearing polyethylene glycol-coupled transferrin", 2003, Neurol. Res. 25:275-279.	
		Orum et al., "Locked nucleic acids: A promising molecular family for gene-function analysis and antisense drug development", 2001, Current Opinion in Molecular Therapeutics 3 (3):239-243.	
		PAN et al., "Isolation of virus-neutralizing RNAs from a large pool of random sequences", 1995, Proc. Natl. Acad. Sci. USA 92:11509-11513.	
		PAPUCCI et al., "Phosphodiester Oligonucleotides Inhibit Mitosis and Trigger Apoptosis by a Non-Antisense, p53-Mediated Mechanism", 2002, Antisense & Nucleic Acid Drug Development 12:21-31.	
		PEREZ et al., "Sequence-independent induction of Sp1 transcription factor activity by phosphorothioate oligodeoxynucleotides", 1994, <i>Proc. Natl. Acad. Sci. USA</i> 91:5957-5961.	
		POLI et al., "In vitro Evaluation of the Anti-prionic Activity of Newly Synthesized Congo Red Derivatives", 2003, ArzneimForsch./Drug Res. 53 (12):875-888.	

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		PRIOLA et al., "Porphyrin and Phthalocyanine Antiscrapie Compounds", 2000, Science 287: 1503-1506.	
		PROSKE <i>et al.</i> , "Prion-Protein-Specific Aptamer Reduces PrP ^{Sc} Formation", 2002, <i>Chemic. Biol.Chem.</i> 3:717-725.	
		RAMAZEILLES et al., "Antisense phosphorothioate oligonucleotides: Selective killing of the intracellular parasite Leishmania amazonensis", 1994, Proc. Natl. Acad. Sci. USA 91:7859-7863.	
		RHIE et al., "Characterization of 2'-fluoro-RNA aptamers that bind preferentially to disease-associated conformations of prion protein and inhibit conversion", 2003, J. Biol. Chem., 278 (41):39697-39705.	
		Rieger, in: <i>Pharmaceutical Dosage Forms</i> , Lieberman, Rieger and Banker (Eds), 1988, Marcel Dekker, INC., New York, NY, Vol. 1, pp. 285-366	
		ROH et al., "Down-Regulation of HER2/neu Expression Induces Apoptosis in Human Cancer Cells That Overexpress HER2/neu", 2000, Cancer Research 60:560-565.	
:	_	ROSOFF, in: <i>Pharmaceutical Dosage Forms</i> , Lieberman, Rieger and Banker (Eds), 1988, Marcel Dekker, INC., New York, NY, Vol. 1, pp.245-282.	
		SCHMIDT et al., "Drug targeting by long-circulating liposomal glucocorticosteroids increases therapeutic efficacy in a model of multiple sclerosis", 2003, <i>Brain</i> 126:1895-1904.	
		SHYNG <i>et al.</i> , "Sulfated Glycans Stimulate Endocytosis of the Cellular Isoform of the Prion Protein, PrP ^c , in Cultured Cells", 1995, <i>J. Biol. Chem.</i> 270 (50):30221-30229.	
		SIERAKOWSKA <i>et al.</i> , "Repair of thalassemic human β-globin mRNA in mammalian cells by antisense oligonucleotides", 1996, <i>Proc. Natl. Acad. Sci. USA</i> 93:12840-12844.	

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		SMETSERS et al., "An antisense Bcr-Abl phosphodiester-tailed methylphosphonate oligonucleotide reduces the growth of chronic myeloid leukaemia patient cells bay a non-antisense mechanism", 1997, British Journal of Haematology 96:377-381.	
		SUPATTAPONE et al., "Branched Polyamines Cure Prion-Infected Neuroblastoma Cells", 2001, J. Virol. 75 (7):3453-3461.	
		TCHATALBACHEV et al., "The packaging signal of influenza viral RNA molecules", 2001, RNA 7: 979-989.	
		UHLMANN and PEYMAN, "Antisense Oligonucleotides: A New Therapeutic Principle", 1990, Chemical Reviews 90 (4):544-584.	
		VAN DEVENTER et al., "A randomised, controlled, double blind, escalating dose study of alicaforsen enema in active ulcerative colitis", 2004, <i>Gut.</i> 53:1646-1651.	
		VINOGRADOV et al., "Nanogels for Oligonucleotide Delivery to the Brain", 2004, Bioconjug. Chem. 15:50-60.	
		WANG et al., "Sequence-independent Inhibition of In Vitro Vascular Smooth Muscle Cell Proliferation, Migration, and In Vivo Neointimal Formation by Phosphorothioate Oligodeoxynucleotides", 1996, J. Clin. Invest. 98 (2):443-450.	
		WHITE et al., "Inhibition of the Multiple Antibiotic Resistance (mar) Operon in Escherichia coli by Antisense DNA Analogs", 1997, Antimicrobial Agents and Chemotherapy 41 (12):2699-2704.	
		WHITE et al., "Development of novel methods for delivering therapeutic oligonucleotides to the central nervous system", 2003, Society for Neuroscience, Program #325.5, Abstract.	
		WHITE et al., "Antisense oligonucleotide treatments for psoriasis", 2004, Expert. Opin. Biol. Ther. 4(1):75-81.	

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		XU et al., "Inhibition of DNA Replication and Induction of S Phase Cell Cycle Arrest by G-rich Oligonucleotides", 2001, The Journal of Biological Chemistry 276 (46):43221-43230.	
		YANG et al., "Construction and selection of bead-bound combinatorial oligonucleoside phosphorothioate and phosphorodithioate aptamer libraries designed for rapid PCR-based sequencing", 2002, <i>Nucl. Acids Res.</i> 30 (23):1-8.	
		YU et al., "Hybrid Oligonucleotides: Synthesis, Biophysical Properties, Stability Studies, and Biological Activity", 1996, Bioorganic. Med. Chem. 4 (10):1685-1692.	
		YU et al., "Prediction of Clinical Responses in a Simulated Phase III Trial of Crohn's Patients Administered the Antisense Phosphorothioate Oligonucleotide ISIS 2302: Comparison of Proposed Dosing Regimens", 2003, Antisense Nucleic Acid Drug Dev. 13:57-66.	
		ZELLWEGER et al., "Antitumor Activity of Antisense Clusterin Oligonucleotides Is Improved in Vitro and in Vivo by Incorporation of 2'-O-(2-Methoxy)Ethyl Chemistry", 2001, J. Pharmacol. and Experimental Therapeutics 298 (3):934-940.	
		ZHANG et al., "A Simple Glycol Nucleic Acid", 2005, J. Am. Chem. Soc. 127:4174-4175.	
		ZHANG et al., "The Study on Brain Targeting of the Amphotericin B Liposomes", 2003, J. Drug. Target. 11 (2):117-122.	
		ZHANG et al., "Global Non-Viral Gene Transfer to the Primate Brain Following Intravenous Administration", 2003, Mol. Ther. 7 (1):11-18.	
		The Concise Encyclopedia of Polymer Science and Engineering, Jacqueline I. Kroschwitz, 1998, ISBN: 0-471-31856-6, 1341 pages, pp. 858-859.	
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